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<110> Bennett, Michele
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      Krause, James
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300

Cys His Leu Thr Ala Met Ile Ser Thr Cys Val Asn Pro Ile Phe Tyr 305 310 315 320

Gly Phe Leu Asn Lys Asn Phe Gln Arg Asp Leu Gln Phe Phe Asn 325 330 335

Phe Cys Asp Phe Arg Ser Arg Asp Asp Asp Tyr Glu Thr Ile Ala Met 340 345 350

Ser Thr Met His Thr Asp Val Ser Lys Thr Ser Leu Lys Gln Ala Ser 355 360 365

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Val Ile Ile Leu Gly Val Ser Gly Asn Leu Ala Leu Ile Ile Ile 50 60

Leu Lys Gln Lys Glu Met Arg Asn Val Thr Asn Ile Leu Ile Val Asn 65 70 75 80

Leu Ser Phe Ser Asp Leu Leu Val Ala Val Met Cys Leu Pro Phe Thr 85 90 95

Phe Val Tyr Thr Leu Met Asp His Trp Val Phe Gly Glu Thr Met Cys 100 105 110

Lys Leu Asn Pro Phe Val Gln Cys Val Ser Ile Thr Val Ser Ile Phe 115 120 125

Ser Leu Val Leu Ile Ala Val Glu Arg His Gln Leu Ile Ile Asn Pro 130 135 140

Arg Gly Trp Arg Pro Asn Asn Arg His Ala Tyr Ile Gly Ile Thr Val 145 150 155 160

Ile Trp Val Leu Ala Val Ala Ser Ser Leu Pro Phe Val Ile Tyr Gln

165 . 170 175

Ile Leu Thr Asp Glu Pro Phe Gln Asn Val Ser Leu Ala Ala Phe Lys 180 185 190

Asp Lys Tyr Val Cys Phe Asp Lys Phe Pro Ser Asp Ser His Arg Leu 195 200 205

Ser Tyr Thr Thr Leu Leu Leu Val Leu Gln Tyr Phe Gly Pro Leu Cys 210 225 220

Phe Ile Phe Ile Cys Tyr Phe Lys Ile Tyr Ile Arg Leu Lys Arg Arg 225 230 235 240

Asn Asn Met Met Asp Lys Ile Arg Asp Ser Lys Tyr Arg Ser Ser Glu 245 250 255

Thr Lys Arg Ile Asn Val Met Leu Leu Ser Ile Val Val Ala Phe Ala 260 265 270

Val Cys Trp Leu Pro Leu Thr Ile Phe Asn Thr Val Phe Asp Trp Asn 275 280 285

His Gln Ile Ile Ala Thr Cys Asn His Asn Leu Leu Phe Leu Leu Cys 290 . 295 300

His Leu Thr Ala Met Ile Ser Thr Cys Val Asn Pro Ile Phe Tyr Gly 305 310 315 320

Phe Leu Asn Lys Asn Phe Gln Arg Asp Leu Gln Phe Phe Asn Phe 325 330 335

Cys Asp Phe Arg Ser Arg Asp Asp Tyr Glu Thr Ile Ala Met Ser 340 345 350

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Val Ala Phe Lys Lys Ile Ser Met Asn Asp Asn Glu Lys Ile 370 375 380

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<212> DNA

<213> Homo sapiens

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tgatttccca gtctgggatg actataaaag cagtgtagat gacttacagt attttctgat 180
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Arg Asn Ser Asp Phe Pro Val Trp Asp Asp Tyr Lys Ser Ser Val Asp 35 40 45

Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu Leu 50 55 60

Gly Phe Met Gly Asn Leu Leu Ile Leu Met Ala Leu Met Lys Lys Arg
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Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe Ser 85 90 95

Asp Ile Leu Val Val Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser Val 100 105 110

Leu Leu Asp Gln Trp Met Phe Gly Lys Val Met Cys His Ile Met Pro 115 120 125

Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile Ser 130 135 140

Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn Asn 145 150 155 160

Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr Leu 165 170 175

Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val Glu 180 185 190

Leu Gln Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Arg Tyr Leu Cys 195 200 205

Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile Ser 210 225 220

Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val Ser 225 230 235 235

His Thr Ser Val Cys Ile Arg Leu Lys Arg Arg Asn Asn Met Met Asp 245 250 255

Lys Met Arg Asp Asn Lys Tyr Arg Ser Ser Arg Ser Arg Ser Val Phe 260 265 270

Tyr Arg Leu Thr Ile Leu Ile Leu Val Phe Ala Val Ser Trp Met Pro 275 280 285

Leu His Leu Phe His Val Val Thr Asp Phe Asn Asp Asn Leu Ile Ser 290 295 300

Asn Arg His Phe Lys Leu Val Tyr Cys Ile Cys His Leu Leu Gly Met 305 310 315 320

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tgatttccca gtctgggatg actataaaag cagtgtagat gacttacagt attttctgat 180
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tgattatgaa acaatagcca tgtccacgat gcacacagat gtttccaaaa cttctttgaa 1140
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<223> Description of Artificial Sequence: Y5/Y1 CHIMERA

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Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu Leu 55 60

Gly Phe Met Gly Asn Leu Leu Ile Leu Met Ala Leu Met Lys Lys Arg 70 65

Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe Ser 90

Asp Ile Leu Val Val Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser Val 100 105 110

Leu Leu Asp Gln Trp Met Phe Gly Lys Val Met Cys His Ile Met Pro 115 120

Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile Ser 135 140

Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn Asn 145 150

Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr Leu Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val Glu Leu Gln Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Arg Tyr Leu Cys 200 Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile Ser Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val Ser 230 235 His Thr Ser Val Cys Arg Ser Ile Ser Cys Gly Leu Ser Asn Lys Glu 250 Asn Arg Leu Glu Glu Asn Glu Met Ile Asn Leu Thr Leu His Pro Ser Lys Lys Ser Gly Pro Gln Val Lys Leu Ser Gly Ser His Lys Trp Ser 275 Tyr Ser Phe Ile Lys Lys His Arg Arg Arg Tyr Ser Lys Lys Thr Ala 295 Cys Val Leu Pro Ala Pro Glu Arg Pro Ser Gln Glu Asn His Ser Arg 315 310 305 Ile Leu Pro Glu Asn Phe Gly Ser Val Arg Ser Gln Leu Ser Ser Ser 325 330 Ser Lys Phe Ile Pro Gly Val Pro Thr Cys Phe Glu Ile Lys Pro Glu 345 340 Glu Asn Ser Asp Val His Glu Leu Arg Val Lys Arg Ser Val Thr Arg 360 365 Ile Lys Lys Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile 375 380 Leu Val Phe Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val 395 Thr Asp Phe Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val Tyr Cys Ile Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr Gly Phe Leu Asn Asn Gly Ile Gln Arg Asp Leu Gln Phe 440 Phe Phe Asn Phe Cys Asp Phe Arg Ser Arg Asp Asp Tyr Glu Thr

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Ile Ala Met Ser Thr Met His Thr Asp Val Ser Lys Thr Ser Leu Lys 465 470 480

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Glu Lys Ile

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<211> 394

<212> PRT

<213> Artificial Sequence

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<223> Description of Artificial Sequence:Y5/Y1 CHIMERA

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Arg Asn Ser Asp Phe Pro Val Trp Asp Asp Tyr Lys Ser Ser Val Asp 35 40 45

Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu Leu 50 60

Gly Phe Met Gly Asn Leu Leu Ile Leu Met Ala Leu Met Lys Lys Arg
65 70 75 80

Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe Ser 85 90 95

Asp Ile Leu Val Val Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser Val

Leu Leu Asp Gln Trp Met Phe Gly Lys Val Met Cys His Ile Met Pro 115 120 125

Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile Ser 130 135 140

Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn Asn 145 150 155 160

Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr Leu 165 170 175

Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val Glu 180 185 190

Leu Gln Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Arg Tyr Leu Cys

195 200 205

Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile Ser 210 225 220

Leu Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val Ser 225 230 235 240

Lys Met Arg Asp Asn Lys Tyr Arg Ser Ser Arg Ser Arg Ser Val Phe 260 265 270

Tyr Arg Leu Thr Ile Leu Ile Leu Val Phe Ala Val Ser Trp Met Pro 275 280 285

Leu His Leu Phe His Val Val Thr Asp Phe Asn Asp Asn Leu Ile Ser 290 295 300

Asn Arg His Phe Lys Leu Val Tyr Cys Ile Cys His Leu Leu Gly Met 305 310 315

Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr Gly Phe Leu Asn Asn Gly 325 330 335

Ile Gln Arg Asp Leu Gln Phe Phe Phe Asn Phe Cys Asp Phe Arg Ser 340 345 350

Arg Asp Asp Tyr Glu Thr Ile Ala Met Ser Thr Met His Thr Asp 355 360 365

Val Ser Lys Thr Ser Leu Lys Gln Ala Ser Pro Val Ala Phe Lys Lys 370 380

Ile Asn Asn Asn Asp Asp Asn Glu Lys Ile 385 390

<210> 11

<211> 6

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: HEXAHISTADINE TAG

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His His His His His 5

<210> 12

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<212> PRT

<213> Artificial Sequence

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Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu Leu
Gly Phe Met Gly Asn Leu Leu Ile Leu Met Ala Leu Met Lys Lys Arg
Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe Ser
Asp Ile Leu Val Val Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser Val
Leu Leu Asp Gln Trp Met Phe Gly Lys Val Met Cys His Ile Met Pro
                            120
Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile Ser
                        135
    130
Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn Asn
                    150
                                        155
Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr Leu
                165
                                    170
Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val Glu
                                185
Leu Gln Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Arg Tyr Leu Cys
                            200
                                                 205
Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile Ser
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Leu Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val Ser 235 His Thr Ser Val Cys Arg Ser Ile Ser Cys Gly Leu Ser Asn Lys Glu Asn Arg Leu Glu Glu Asn Glu Met Ile Asn Leu Thr Leu His Pro Ser 265 Lys Lys Ser Gly Pro Gln Val Lys Leu Ser Gly Ser His Lys Trp Ser Tyr Ser Phe Ile Lys Lys His Arg Arg Tyr Ser Lys Lys Thr Ala 295 Cys Val Leu Pro Ala Pro Glu Arg Pro Ser Gln Glu Asn His Ser Arg 315 Ile Leu Pro Glu Asn Phe Gly Ser Val Arg Ser Gln Leu Ser Ser Ser 330 Ser Lys Phe Ile Pro Gly Val Pro Thr Cys Phe Glu Ile Lys Pro Glu Glu Asn Ser Asp Val His Glu Leu Arg Val Lys Arg Ser Val Thr Arg Ile Lys Lys Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile 370 Leu Val Phe Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val 395 Thr Asp Phe Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val 405 410 Tyr Cys Ile Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro 425 420 Ile Leu Tyr Gly Phe Leu Asn Asn Gly Ile Lys Ala Asp Leu Val Ser 440 445 435

<210> 14 <211> 21 <212> DNA <213> Homo sapiens

Leu Ile His Cys Leu His Met

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450

<211> <212> <213>		
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<210><211><211><212><213>	63	
<220> <223>	Description of Artificial Sequence: CHIMERIC Y1/Y5 PRIMER	
<400> tacgco gta	16 ctaaa aaggagaaac aacatgatgg acaagatgag agacaataag tacaggtcca	60 63
<210> <211> <212> <213>	71	
<220> <223>	Description of Artificial Sequence: CHIMERIC Y1/Y5 PRIMER	
_	17 actgg acctgtactt attgtctctc atcttgtcca tcatgttgtt tctccttttt tatgc a	60 71
<210><211><211><212><213>	31	
<220> <223>	Description of Artificial Sequence: MUTAGENIC R1 PRIMER	
<400> gaacaa	18 aaaga attcagagag acttgcagtt c	31
<210><211><211><212><213>	28	
<220> <223>	Description of Artificial Sequence: MUTAGENIC R1 PRIMER	

<210> 20

<211> 341 <212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Y1/Y5 CHIMERA

<400> 20

Met Asp Leu Glu Leu Gln Asp Phe Tyr Asn Lys Thr Leu Ala Thr Glu

Asn Asn Thr Ala Ala Thr Arg Asn Ser Asp Phe Pro Val Trp Asp Asp 25

Tyr Lys Ser Ser Val Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr

Thr Phe Val Ser Leu Leu Gly Phe Met Gly Asn Leu Leu Ile Leu Met

Ala Leu Met Arg Lys Arg Asn Gln Lys Thr Met Val Asn Phe Leu Ile

Gly Asn Leu Ala Phe Ser Asp Ile Leu Val Val Leu Phe Cys Ser Pro

Phe Thr Leu Thr Ser Val Leu Leu Asp Gln Trp Met Phe Gly Lys Val 110 100

Met Cys His Ile Met Pro Phe Leu Gln Cys Val Ser Val Leu Val Ser 120 115

Thr Leu Ile Leu Ile Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys 140 130 135

His Pro Ile Ser Asn Asn Leu Thr Ala Asn His Gly Tyr Phe Leu Ile 150 155

Ala Thr Val Trp Thr Leu Gly Phe Ala Ile Cys Ser Pro Leu Pro Val 165 170 175

Phe His Ser Leu Val Glu Leu Gln Glu Thr Phe Asp Ser Ala Leu Leu 185

Ser Ser Arg Tyr Leu Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg 200 205

Ile Ala Phe Thr Ile Ser Leu Leu Leu Val Gln Tyr Ile Leu Pro Leu

Val Cys Leu Thr Val Ser His Thr Ser Val Cys Ile Arg Leu Lys Arg

<u>L</u>

Arg Asn Asn Met Met Asp Lys Met Arg Asp Asn Lys Tyr Arg Ser Ser 245 250 255

Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile Leu Val Phe 260 265 270

Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val Thr Asp Phe 275 280 285

Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val Tyr Cys Ile 290 295 300

Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr 305 310 315 320

Gly Phe Leu Asn Asn Gly Ile Lys Ala Asp Leu Ile Ser Leu Ile Gln 325 330 335

Cys Leu His Met Ser 340

<210> 21

<211> 383

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Y1/Y5 CHIMERA

<400> 21

Met Asp Leu Glu Leu Gln Asp Phe Tyr Asn Lys Thr Leu Ala Thr Glu $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$

Asn Asn Thr Ala Ala Thr Arg Asn Ser Asp Phe Pro Val Trp Asp Asp 20 25 30

Tyr Lys Ser Ser Val Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr 35 40 45

Thr Phe Val Ser Leu Leu Gly Phe Met Gly Asn Leu Leu Ile Leu Met 50 55 60

Ala Leu Met Arg Lys Arg Asn Gln Lys Thr Met Val Asn Phe Leu Ile 65 70 . 75 80

Gly Asn Leu Ala Phe Ser Asp Ile Leu Val Val Leu Phe Cys Ser Pro 85 90 95

Phe Thr Leu Thr Ser Val Leu Leu Asp Gln Trp Met Phe Gly Lys Val 100 105 110

Met Cys His Ile Met Pro Phe Leu Gln Cys Val Ser Val Leu Val Ser 115 120 125 Thr Leu Ile Leu Ile Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys 130. 135

His Pro Ile Ser Asn Asn Leu Thr Ala Asn His Gly Tyr Phe Leu Ile 145 150 155 160

Ala Thr Val Trp Thr Leu Gly Phe Ala Ile Cys Ser Pro Leu Pro Val 165 170 175

Phe His Ser Leu Val Glu Leu Gln Glu Thr Phe Asp Ser Ala Leu Leu 180 185 190

Ser Ser Arg Tyr Leu Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg 195 200 205

Ile Ala Phe Thr Ile Ser Leu Leu Leu Val Gln Tyr Ile Leu Pro Leu 210 215 220

Val Cys Leu Thr Val Ser His Thr Ser Val Cys Ile Arg Leu Lys Arg 225 230 235 240

Arg Asn Asn Met Met Asp Lys Met Arg Asp Asn Lys Tyr Arg Ser Ser 245 250 255

Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile Leu Val Phe 260 265 270

Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val Thr Asp Phe 275 280 285

Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val Tyr Cys Ile 290 295 300

Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr 305 310 315 320

Gly Phe Leu Asn Asn Gly Ile Gln Arg Asp Leu Gln Phe Phe Asn 325 330 335

Phe Cys Asp Phe Arg Ser Arg Asp Asp Tyr Glu Thr Ile Ala Met 340 345 350

Ser Thr Met His Thr Asp Val Ser Lys Thr Ser Leu Lys Gln Ala Ser 355 360 365

Pro Val Ala Phe Lys Lys Ile Asn Asn Asp Asp Asn Glu Lys Ile 370 375 380

<210> 22

<211> 508

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Y1/Y5 CHIMERA

<400> 22 Met Glu Val Lys Leu Glu Glu His Phe Asn Lys Thr Phe Val Thr Glu Asn Asn Thr Ala Ala Ser Gln Asn Thr Ala Ser Pro Ala Trp Glu Asp 25 Tyr Arg Gly Thr Glu Asn Asn Thr Ser Ala Ala Arg Asn Thr Ala Phe Pro Val Trp Glu Asp Tyr Arg Gly Ser Val Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu Leu Gly Phe Met Gly Asn 70 Leu Leu Ile Leu Met Ala Val Met Lys Lys Arg Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe Ser Asp Ile Leu Val Val 105 Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser Val Leu Leu Asp Gln Trp 115 Met Phe Gly Lys Ala Met Cys His Ile Met Pro Phe Leu Gln Cys Val 135 Ser Val Leu Val Ser Thr Leu Ile Leu Ile Ser Ile Ala Ile Val Arg 145 Tyr His Met Ile Lys His Pro Ile Ser Asn Asn Leu Thr Ala Asn His 165 170 Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr Leu Gly Phe Ala Ile Cys 185 180 Ser Pro Phe Pro Val Phe His Ser Leu Val Glu Leu Lys Glu Thr Phe 200 Gly Ser Ala Leu Leu Ser Ser Lys Tyr Leu Cys Val Glu Ser Trp Pro 220 215 210 Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile Ser Leu Leu Leu Val Gln 230 235 Tyr Ile Leu Pro Leu Val Cys Leu Thr Val Ser His Thr Ser Val Cys 245 250 255 Arg Ser Ile Ser Cys Gly Leu Ser His Lys Glu Asn Arg Leu Glu Glu 265 Asn Glu Met Ile Asn Leu Thr Leu His Pro Ser Lys Lys Ser Arg Asp 280 Gln Ala Lys Pro Pro Ser Thr Gln Lys Trp Ser Tyr Ser Phe Ile Arg

Lys His Arg Arg Arg Tyr Ser Lys Lys Thr Ala Cys Val Leu Pro Ala Pro Ala Gly Pro Ser Gln Glu Lys His Leu Thr Val Pro Glu Asn Pro Gly Ser Val Arg Ser Gln Leu Ser Pro Ser Ser Lys Val Ile Pro Gly 345 Val Pro Ile Cys Phe Glu Val Lys Pro Glu Glu Ser Ser Asp Ala Gln Glu Met Arg Val Lys Arg Ser Leu Thr Arg Ile Lys Lys Arg Ser Arg 375 Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile Leu Val Phe Ala Val Ser Trp Met Pro Leu His Val Phe His Val Val Thr Asp Phe Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val Tyr Cys Ile Cys His Leu 420 Leu Gly Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr Gly Phe Leu 440 Asn Asn Gly Ile Gln Arg Asp Leu Gln Phe Phe Asn Phe Cys Asp 450 455 Phe Arg Ser Arg Asp Asp Tyr Glu Thr Ile Ala Met Ser Thr Met 470 475 His Thr Asp Val Ser Lys Thr Ser Leu Lys Gln Ala Ser Pro Val Ala 485 490

Phe Lys Lys Ile Ser Met Asn Asp Asn Glu Lys Val
500 505

<210> 23

<211> 352

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Y1/Y5 CHIMERA

<400> 23

Met Asp Val Leu Phe Phe His Gln Asp Ser Ser Met Glu Phe Lys Leu
1 5 10 15

Glu Glu His Phe Asn Lys Thr Phe Val Thr Glu Asn Asn Thr Ala Ala
20 25 30

Ala Arg Asn Ala Ala Phe Pro Ala Trp Glu Asp Tyr Arg Gly Ser Val

35 40 45

Asp	Asp 50	Leu	Gln	Tyr	Phe	Leu 55	Ile	Gly	Leu	Tyr	Thr 60	Phe	Val	Ser	Leu
Leu 65	Gly	Phe	Met	Gly	Asn 70	Leu	Leu	Ile	Leu	Met 75	Ala	Val	Met	Lys	Lys 80
Arg	Asn	Gln	Lys	Thr 85	Thr	Val	Asn	Phe	Leu 90	Ile	Gly	Asn	Leu	Ala 95	Phe ·
Ser	Asp	Ile	Leu 100	Val	Val	Leu	Phe	Cys 105	Ser	Pro	Phe	Thr	Leu 110	Thr	Ser
Val	Leu	Leu 115	Asp	Gln	Trp	Met	Phe 120	Gly	Lys	Ala	Met	Cys 125	His	Ile	Met
Pro	Phe 130	Leu	Gln	Cys	Val	Ser 135	Val	Leu	Val	Ser	Thr 140	Leu	Ile	Leu	Ile
Ser 145	Ile	Ala	Ile	Val	Arg 150	Tyr	His	Met	Ile	Lys 155	His	Pro	Ile	Ser	Asn 160
Asn	Leu	Thr	Ala	Asn 165	His	Gly	Tyr	Phe	Leu 170	Ile	Ala	Thr	Val	Trp 175	Thr
	_		Ala 180					185					190		
Glu	Leu	Lys 195	Glu	Thr	Phe	Gly	Ser 200	Ala	Leu	Leu	Ser	Ser 205	Lys	Tyr	Leu
_	210		Ser	-		215					220				
Ser 225	Leu	Leu	Leu	Val	Gln 230	Tyr	Ile	Leu	Pro	Leu 235	Val	Cys	Leu	Thr	Val 240
Ser	His	Thr	Ser	Val 245	Cys	Ile	Arg	Leu	Lys 250	Arg	Arg	Asn	Asn	Met 255	Met
Asp	Lys	Ile	Arg 260	Asp	Ser	Lys	Tyr	Arg 265	Ser	Ser	Arg	Ser	Arg 270	Ser	Val
Phe	Tyr	Arg 275	Leu	Thr	Ile	Leu	Ile 280	Leu	Val	Phe	Ala	Val 285	Ser	Trp	Met
Pro	Leu 290	His	Val	Phe	His	Val 295	Val	Thr	Asp	Phe	Asn 300	Asp	Asn	Leu	Ile
Ser 305	Asn	Arg	His	Phe	Lys 310	Leu	Val	Tyr	Cys	11e 315	Суѕ	His	Leu	Leu	Gly 320
Met	Met	Ser	Cys	Cys 325	Leu	Asn	Pro	Ile	Leu 330	Tyr	Gly	Phe	Leu	Asn 335.	Asn
Gly	Ile	Lys	Ala	Asp	Leu	Arg	Ala	Leu	Ile	His	Cys	Leu	His	Met	Ser

<210> 24 <211> 499 <212> PRT <213> Artificial Sequence <223> Description of Artificial Sequence:Y1/Y5 CHIMERA <400> 24 Met Asp Val Leu Phe Phe His Gln Asp Ser Ser Met Glu Phe Lys Leu Glu Glu His Phe Asn Lys Thr Phe Val Thr Glu Asn Asn Thr Ala Ala 25 Ala Arg Asn Ala Ala Phe Pro Ala Trp Glu Asp Tyr Arg Gly Ser Val Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu Leu Gly Phe Met Gly Asn Leu Leu Ile Leu Met Ala Val Met Lys Lys Arg Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe Ser Asp Ile Leu Val Val Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser 105 Val Leu Leu Asp Gln Trp Met Phe Gly Lys Ala Met Cys His Ile Met 120 125 115 Pro Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile 135 Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn 150 155 145 Asn Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr 170 165 Leu Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val 180 185 190 Glu Leu Lys Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Lys Tyr Leu 200

Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile

220

215

Ser Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val 235 230 Ser His Thr Ser Val Cys Arg Ser Ile Ser Cys Gly Leu Ser His Lys 245 250 Glu Asn Arg Leu Glu Glu Asn Glu Met Ile Asn Leu Thr Leu Gln Pro 265 Ser Lys Lys Ser Arg Asn Gln Ala Lys Thr Pro Ser Thr Gln Lys Trp 280 Ser Tyr Ser Phe Ile Arg Lys His Arg Arg Arg Tyr Ser Lys Lys Thr Ala Cys Val Leu Pro Ala Pro Ala Gly Pro Ser Gln Gly Lys His Leu 315 310 Ala Val Pro Glu Asn Pro Ala Ser Val Arg Ser Gln Leu Ser Pro Ser Ser Lys Val Ile Pro Gly Val Pro Ile Cys Phe Glu Val Lys Pro Glu 345 Glu Ser Ser Asp Ala His Glu Met Arg Val Lys Arg Ser Ile Thr Arg Ile Lys Lys Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile 375 Leu Val Phe Ala Val Ser Trp Met Pro Leu His Val Phe His Val Val 385 Thr Asp Phe Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val 410 Tyr Cys Ile Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro 425 420 Ile Leu Tyr Gly Phe Leu Asn Asn Gly Ile Lys Gln Arg Asp Leu Gln 440 Phe Phe Phe Asn Phe Cys Asp Phe Arg Ser Arg Asp Asp Asp Tyr Glu 450 455 Thr Ile Ala Met Ser Thr Met His Thr Asp Val Ser Lys Thr Ser Leu 475 470 Lys Gln Ala Ser Pro Val Ala Phe Lys Lys Ile Ser Met Asn Asp Asn 490 485

<210> 25 <211> 395

Glu Lys Ile

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Y1/Y5 CHIMERA

<400> 25

Met Asp Val Leu Phe Phe His Gln Asp Ser Ser Met Glu Phe Lys Leu
1 5 10 15

Glu Glu His Phe Asn Lys Thr Phe Val Thr Glu Asn Asn Thr Ala Ala
20 25 30

Ala Arg Asn Ala Ala Phe Pro Ala Trp Glu Asp Tyr Arg Gly Ser Val\$35\$ 40 45

Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr Thr Phe Val Ser Leu 50 60

Leu Gly Phe Met Gly Asn Leu Leu Ile Leu Met Ala Val Met Lys Lys 65 70 75 80

Arg Asn Gln Lys Thr Thr Val Asn Phe Leu Ile Gly Asn Leu Ala Phe 85 90 95

Ser Asp Ile Leu Val Val Leu Phe Cys Ser Pro Phe Thr Leu Thr Ser 100 105 110

Val Leu Leu Asp Gln Trp Met Phe Gly Lys Ala Met Cys His Ile Met 115 120 125

Pro Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile 130 135 140

Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn 145 150 155 160

Asn Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr 165 170 175

Leu Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val 180 185 190

Glu Leu Lys Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Lys Tyr Leu 195 200 205

Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile 210 215 · 220

Ser Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val 225 230 235 240

Ser His Thr Ser Val Cys Ile Arg Leu Lys Arg Arg Asn Asn Met Met 245 250 255

Asp Lys Ile Arg Asp Ser Lys Tyr Arg Ser Ser Arg Ser Arg Ser Val 260 265 270

Phe Tyr Arg Leu Thr Ile Leu Ile Leu Val Phe Ala Val Ser Trp Met 275 280 285

Pro Leu His Val Phe His Val Val Thr Asp Phe Asn Asp Asn Leu Ile 290 295 300

Ser Aşn Arg His Phe Lys Leu Val Tyr Cys Ile Cys His Leu Leu Gly 305 310 315 320

Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr Gly Phe Leu Asn Asn 325 330 335

Gly Ile Lys Gln Arg Asp Leu Gln Phe Phe Phe Asn Phe Cys Asp Phe 340 345 350

Arg Ser Arg Asp Asp Asp Tyr Glu Thr Ile Ala Met Ser Thr Met His 355 360 365 .

Thr Asp Val Ser Lys Thr Ser Leu Lys Gln Ala Ser Pro Val Ala Phe 370 375 380

Lys Lys Ile Ser Met Asn Asp Asn Glu Lys Ile 385 390 395

<210> 26

<211> 341

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:Y1/Y5 CHIMERA

<400> 26

Met Gly Ser Glu Ile Pro Asp Tyr Tyr Asn Lys Thr Leu Ala Ser Glu 1 5 10 15

Asn Asn Thr Val Ala Thr Arg Asn Ser Gly Phe Pro Val Trp Glu Asp
20 25 30 .

Tyr Lys Gly Ser Val Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr 35 40 45

Thr Phe Val Ser Leu Leu Gly Phe Met Gly Asn Leu Leu Ile Leu Met 50 55 60

Ala Val Met Arg Lys Arg Asn Gln Lys Thr Thr Val Asn Phe Leu Ile
65 70 75 80

Gly Asn Leu Ala Phe Ser Asp Ile Leu Val Val Leu Phe Cys Ser Pro 85 90 95

Phe Thr Leu Thr Ser Val Leu Leu Asp Gln Trp Met Phe Gly Lys Val 100 105 110

Met Cys His Ile Met Pro Phe Leu Gln Cys Val Thr Val Leu Val Ser

115 120 125

Thr Leu Ile Leu Ile Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys
130 140

His Pro Val Ser Asn Asn Leu Thr Ala Asn His Gly Tyr Phe Leu Ile 145 150 155 160

Ala Thr Val Trp Thr Leu Gly Leu Ala Ile Cys Ser Pro Leu Pro Val 165 170 175

Phe His Ser Leu Val Glu Leu Gln Glu Ser Phe Gly Ser Ala Trp Leu 180 185 190

Ser Ser Arg Tyr Leu Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg 195 200 205

Ile Ala Phe Thr Ile Ser Leu Leu Leu Val Gln Tyr Ile Leu Pro Leu 210 215 220

Val Cys Leu Thr Val Ser His Thr Ser Val Cys Ile Arg Leu Lys Arg 225 230 235 240

Arg Asn Asn Met Met Asp Lys Met Arg Asp Asn Lys Tyr Arg Ser Ser 245 250 255

Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Val Leu Ile Leu Val Phe 260 265 270

Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val Thr Asp Phe 275 280 285

Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val Tyr Cys Ile 290 295 300

Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr 305 310 315 320

Gly Phe Leu Asn Asn Gly Ile Lys Ala Asp Leu Met Ser Leu Ile His 325 330 335

Cys Leu His Val Ser 340

<210> 27

<211> 383

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Y1/Y5 CHIMERA

<400> 27

Met Gly Ser Glu Ile Pro Asp Tyr Tyr Asn Lys Thr Leu Ala Ser Glu 1 5 10 15

Asn Asn Thr Val Ala Thr Arg Asn Ser Gly Phe Pro Val Trp Glu Asp 20 25 30

Tyr Lys Gly Ser Val Asp Asp Leu Gln Tyr Phe Leu Ile Gly Leu Tyr 35 40 45.

Thr Phe Val Ser Leu Leu Gly Phe Met Gly Asn Leu Leu Ile Leu Met 50 55 60

Ala Val Met Arg Lys Arg Asn Gln Lys Thr Thr Val Asn Phe Leu Ile 65 70 75 80

Gly Asn Leu Ala Phe Ser Asp Ile Leu Val Val Leu Phe Cys Ser Pro 85 90 95

Phe Thr Leu Thr Ser Val Leu Leu Asp Gln Trp Met Phe Gly Lys Val 100 105 110

Met Cys His Ile Met Pro Phe Leu Gln Cys Val Thr Val Leu Val Ser 115 120 125

Thr Leu Ile Leu Ile Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys 130 135 140

His Pro Val Ser Asn Asn Leu Thr Ala Asn His Gly Tyr Phe Leu Ile 145 150 155 160

Ala Thr Val Trp Thr Leu Gly Leu Ala Ile Cys Ser Pro Leu Pro Val 165 170 175

Phe His Ser Leu Val Glu Leu Gln Glu Ser Phe Gly Ser Ala Trp Leu 180 185 190

Ser Ser Arg Tyr Leu Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg 195 200 205

Ile Ala Phe Thr Ile Ser Leu Leu Leu Val Gln Tyr Ile Leu Pro Leu 210 215 220

Val Cys Leu Thr Val Ser His Thr Ser Val Cys Ile Arg Leu Lys Arg 225 230 235 240

Arg Asn Asn Met Met Asp Lys Met Arg Asp Asn Lys Tyr Arg Ser Ser 245 250 255

Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Val Leu Ile Leu Val Phe 260 265 270

Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val Thr Asp Phe 275 280 285

Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val Tyr Cys Ile 290 295 300

Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro Ile Leu Tyr 305 310 315 320

Gly	Phe	Leu	Asn	Asn 325	Gly	Ile	Gln	Arg	Asp 330	Leu	Gln	Phe	Phe	Phe 335	Asn	
Phe	Cys	Asp	Phe 340	Arg	Ser	Arg	Asp	Asp 345	Asp	Tyr	Glu	Val	Ile 350	Ala	Met .	
Ser	Thr	Met 355	His	Thr	Asp	Val	Ser 360	Lys	Thr	Ser	Leu	Lys 365	Gln	Ala	Ser	
Pro	Val 370	Ala	Leu	Lys	Lys	Ile 375	His	Ser	Asp	Asp	Asn 380	Glu	Lys	Ile		
<210> 28 <211> 21 <212> DNA <213> Homo sapiens																
)> 28 :ggtt		igaca	aaatq	gt c											21
<210> 29 <211> 26 <212> DNA <213> Homo sapiens																
<400> 29 ccttggtaaa cagtgagaat tattac 26													26			
<210> 30 <211> 455 <212> PRT <213> Cercopithecus aethiops																
)> 3(Ser		Tyr	Ser 5	Lys	Gln	Asp	Tyr	Asn 10	Met	Asp	Leu	Glu	Leu 15	Asp	
Glu	Tyr	Tyr	Asn 20	Lys	Thr	Leu	Ala	Thr 25	Glu	Asn	Asn	Thr	Ala 30	Ala	Thr	
Arg	Asn	Ser 35	Asp	Phe	Pro	Val	Trp 40	Asp	Asp	Tyr	Lys	Ser 45	Ser	Val	Asp	
Asp	Leu 50	Gln	Tyr	Phe	Leu	Ile 55	Gly	Leu	Tyr	Thr	Phe 60	Val	Ser	Leu	Leu	
Gly 65	Phe	Met	Gly	Asn	Leu 70	Leu	Ile	Leu	Met	Ala 75	Leu	Met	Lys	Lys	Arg 80	
Asn	Gln	Lys	Thr	Thr 85	Val	Asn	Phe	Leu	Ile 90	Gly	Asn	Leu	Ala	Phe 95	Ser	
Asp	Ile	Leu	Val 100	Val	Leu	Phe	Суѕ	Ser 105	Pro	Phe	Thr	Leu	Thr 110	Ser	Val	

Leu Leu Asp Gln Trp Met Phe Gly Lys Val Met Cys His Ile Met Pro Phe Leu Gln Cys Val Ser Val Leu Val Ser Thr Leu Ile Leu Ile Ser Ile Ala Ile Val Arg Tyr His Met Ile Lys His Pro Ile Ser Asn Asn 150 155 Leu Thr Ala Asn His Gly Tyr Phe Leu Ile Ala Thr Val Trp Thr Leu 170 Gly Phe Ala Ile Cys Ser Pro Leu Pro Val Phe His Ser Leu Val Glu 185 Leu Gln Glu Thr Phe Gly Ser Ala Leu Leu Ser Ser Arg Tyr Leu Cys Val Glu Ser Trp Pro Ser Asp Ser Tyr Arg Ile Ala Phe Thr Ile Ser 215 Leu Leu Val Gln Tyr Ile Leu Pro Leu Val Cys Leu Thr Val Ser His Thr Ser Val Cys Arg Ser Ile Ser Cys Gly Leu Ser Asn Lys Glu Asn Arg Leu Glu Glu Asn Glu Met Ile Asn Leu Thr Leu His Pro Ser 265 260 Arg Lys Ile Gly Pro Gln Val Lys Leu Ser Gly Ser His Lys Trp Ser 280 Tyr Ser Phe Ile Lys Lys His Arg Arg Tyr Ser Lys Lys Thr Ala 290 295 Cys Val Leu Pro Ala Pro Glu Arg Pro Ser Gln Glu Asn His Ser Arg 310 315 Ile Leu Pro Glu Asn Phe Gly Ser Val Arg Ser Gln Leu Ser Ser Ser 325 330 Ser Lys Phe Ile Pro Gly Val Pro Thr Cys Phe Glu Ile Lys Pro Glu 345 Glu Asn Ser Asp Val His Glu Leu Arg Val Lys Arg Ser Val Thr Arg 365 Ile Lys Lys Arg Ser Arg Ser Val Phe Tyr Arg Leu Thr Ile Leu Ile 375 Leu Val Phe Ala Val Ser Trp Met Pro Leu His Leu Phe His Val Val 395 Thr Asp Phe Asn Asp Asn Leu Ile Ser Asn Arg His Phe Lys Leu Val 405 410

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Tyr Cys Ile Cys His Leu Leu Gly Met Met Ser Cys Cys Leu Asn Pro
420 425 430
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Ile Leu Tyr Gly Phe Leu Asn Asn Gly Ile Lys Ala Asp Leu Met Ser 435 440 445

Leu Ile His Cys Leu His Met 450 455

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